

NDA 18-052/S-025

AUG 11 1998

Schering-Plough HealthCare Products
Attn: Ronald J. Garutti, M.D., Vice President
Clinical Research/Regulatory Affairs
110 Allen Road
P.O. Box 276
Liberty Corner, NJ 07938-0276

Dear Dr. Garutti:

Please refer to your supplemental new drug application dated February 12, 1998, received February 12, 1998, submitted under section 505(b) of the Federal Food, Drug, and Cosmetic Act for Nonprescription Gyne-Lotrimin Vaginal Cream.

We acknowledge receipt of your communications dated April 24, July 23 and 29, and August 10, 1998.

The user fee goal date for this supplemental new drug application is August 12, 1998.

We note that this supplemental NDA was submitted as "Special Supplement-Changes Being Effected" under 21 CFR 314.70(c).

This supplemental new drug application provides for a change in the tamper-resistant packaging for the drug product.

We have completed the review of this supplemental new drug application and have concluded that adequate information has been presented to demonstrate that the drug product is safe and effective for use as recommended in the draft labeling dated February 12, 1998. Accordingly the supplemental new drug application is approved effective on the date of this letter.

Please submit 20 copies of the FPL as soon as it is available, in no case more than 30 days after it is printed. Please individually mount ten of the copies on heavy-weight paper or similar material. For administrative purposes, this submission should be designated "FPL for approved supplemental NDA 18-052/S-025 ." Approval of this submission by FDA is not required before the labeling is used.

As stated in your letters dated July 23 and August 10, 1998, you agreed to revise the labeling for this drug product at the time of the next printing or within 180 days, whichever comes first, as follows:

1. The statement of identity will be revised to read:

“Clotrimazole Vaginal Cream (1%)
Vaginal Antifungal”
2. The tradename “Gyne-Lotrimin 7” will be used in all portions of the labeling for consistency.
3. The statement “Do not purchase if carton is opened” will be deleted from the top panel, since the carton is not part of the tamper-resistant feature, but the term “safety sealed” will be retained as part of the tamper-resistant feature.
4. The text on the back panel will read “...use this cream for seven days in a row.”

If additional information relating to the safety or effectiveness of this drug becomes available before we receive the final printed labeling, revision of that labeling may be required.

Please submit three copies of the introductory promotional materials that you propose to use for this product. All proposed materials should be submitted in draft or mock-up form, not final print. Please submit one copy to this Division and two copies of both the promotional materials and the package insert directly to:

Division of Drug Marketing, Advertising, and Communications
HFD40
Food and Drug Administration
5600 Fishers Lane
Rockville, MD 20857

If a letter communicating important information about this drug product (i.e., a “Dear Healthcare Practitioner” letter) is issued to physicians and others responsible for patient care, we request that you submit a copy of the letter to this NDA and a copy to the following address:

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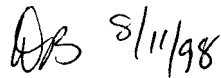
MEDWATCH, HF-2
Food and Drug Administration
5600 Fishers Lane
Rockville, MD 20857

Please submit one market package of the drug product when it is available.

We remind you that you must comply with the requirements for an approved NDA set forth under 21 CFR 314.80 and 314.81

If you have any questions, please contact Sakineh Walther, Project Manager, at (301) 827-2222.

Sincerely yours,

Handwritten signature of Debra L. Bowen, dated 8/11/98.

Debra L. Bowen, M.D.
Acting Director
Division of Over-the-Counter Drug Products
Office of Drug Evaluation V
Center for Drug Evaluation and Research

APPROVED

AUG 11 1998

Gyne-Lotrimin 7-Day Treatment
ANTIFUNGAL Clotrimazole
Vaginal Cream

Cures most vaginal yeast infections
Gyne-Lotrimin 7-Day Treatment
ANTIFUNGAL Clotrimazole
Vaginal Cream

Safety Sealed: Do not purchase if carton is opened. The tube opening should be sealed. If seal has been punctured or the embossed design SP is not visible, do not use product.
Active Ingredient: Clotrimazole 1%
Inactive Ingredients: Benzyl alcohol, cetearyl alcohol, cetyl esters wax, octyldodecanol, polysorbate 60, purified water, sorbitan monostearate

Cures most vaginal yeast infections

Gyne-Lotrimin 7-Day Treatment
ANTIFUNGAL Clotrimazole
Vaginal Cream

EDUCATIONAL PAMPHLET ENCLOSED

ONE 45g (1.5 oz) TUBE OF VAGINAL CREAM (1%) AND APPLICATOR (7-Day Therapy)



Cures most vaginal yeast infections

Gyne-Lotrimin 7-Day Treatment
ANTIFUNGAL Clotrimazole
Vaginal Cream

R-97-34



Store at room temperature between 2° and 30°C (36° and 86°F).

your doctor. You could be pregnant or there could be a serious underlying medical cause for your infection, including diabetes or a damaged immune system (including damage from infection with HIV—the virus that causes AIDS). (PLEASE READ ENCLOSED EDUCATIONAL PAMPHLET). Do not use during pregnancy except under the advice and supervision of a doctor. Do not use tampons while using this medication. Keep this and all drugs out of the reach of children. In case of accidental ingestion, seek professional assistance or contact a Poison Control Center immediately. NOT FOR USE IN CHILDREN LESS THAN 12 YEARS OF AGE.
If you have any questions about GYNE-LOTTRIMIN 7 or vaginal yeast infection, contact your doctor.

See end panel of carton and tube crimp for lot number and expiration date.
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Made in USA
2042403 Schering-Plough

GYNE-LOTTRIMIN 7 Antifungal Vaginal Cream can kill the yeast that may cause vaginal infection. It is greaseless and does not stain clothes.
INDICATIONS: For the treatment of vaginal yeast (Candida) infection and the relief of external vulvar itching and irritation associated with a yeast infection. IF THIS IS THE FIRST TIME YOU HAVE HAD VAGINAL OR VULVAR ITCH AND DISCOMFORT, CONSULT YOUR DOCTOR. IF YOU HAVE HAD A DOCTOR DIAGNOSE A VAGINAL YEAST INFECTION BEFORE AND HAVE THE SAME SYMPTOMS NOW, USE THIS CREAM AS DIRECTED.
WARNING: DO NOT USE IF YOU HAVE ABDOMINAL PAIN, FEVER, OR FOUR-SMELLING DISCHARGE. CONTACT YOUR DOCTOR IMMEDIATELY.
Before using, read the enclosed pamphlet.
Directions: Fill the applicator and insert one applicatorful of cream into the vagina, preferably at bedtime. Repeat this procedure daily for 7 consecutive days. For relief of external vulvar itching, squeeze a small amount of cream onto your finger and gently spread the cream onto the irritated area of the vulva. Use once or twice a day for up to 7 days as needed to relieve external vulvar itching. THE CREAM SHOULD NOT BE USED FOR VULVAR ITCHING DUE TO CAUSES OTHER THAN A YEAST INFECTION.
WARNING: IF YOUR SYMPTOMS DO NOT IMPROVE IN 3 DAYS OR IF YOUR INFECTION ISN'T GONE IN 7 DAYS, YOU MAY HAVE A CONDITION OTHER THAN A YEAST INFECTION. CONSULT YOUR DOCTOR. If your symptoms return within two months or if you have infections that do not clear up easily with proper treatment, consult

Gyne-Lotrimin 7-Day Treatment
ANTIFUNGAL Clotrimazole
Vaginal Cream

Cures most vaginal yeast infections

NO INK CODE AREA NO VARNISH

Cures most vaginal yeast infections
Gyne-Lotrimin 7-Day Treatment
ANTIFUNGAL Clotrimazole
Vaginal Cream

NO VARNISH

Gyne-Lotrimin 7-Day Treatment
ANTIFUNGAL Clotrimazole
Vaginal Cream

Cures most vaginal yeast infections

Safety Sealed. Do not use if seal over tube opening has been punctured or the embossed design **SP** is not visible.

Gyne-Lotrimin[®]

Clotrimazole vaginal cream 1%
ANTIFUNGAL



Net Weight 45g (1.5 oz)



FOR VAGINAL USE ONLY. DO NOT TAKE BY MOUTH OR USE IN EYES

To Open: Use cap to puncture seal.

IF THIS IS THE FIRST TIME YOU HAVE HAD VAGINAL OR VULVAR ITCH AND DISCOMFORT, CONSULT YOUR DOCTOR. IF YOU HAVE HAD A DOCTOR DIAGNOSE A VAGINAL YEAST INFECTION BEFORE AND HAVE THE SAME SYMPTOMS NOW, USE THIS CREAM AS DIRECTED.

GYNE-LOTRIMIN can kill the yeast that may cause vaginal infection.

Greaseless formula that will not stain clothes.

WARNING: DO NOT USE IF YOU HAVE ABDOMINAL PAIN, FEVER, OR FOUL-SMELLING DISCHARGE. CONTACT YOUR DOCTOR IMMEDIATELY.

Before using, read the enclosed pamphlet.

Directions: Fill the applicator and insert one applicatorful of cream into the vagina, preferably at bedtime. Repeat this procedure daily for 7 consecutive days. For relief of external vulvar itching, squeeze a small amount of cream onto your finger and gently spread the cream onto the irritated area of the vulva. Use once or twice a day for up to 7 days as needed to relieve external vulvar itching.

WARNING: IF YOUR SYMPTOMS DO NOT IMPROVE IN 3 DAYS OR IF YOUR INFECTION ISN'T GONE IN 7 DAYS, YOU MAY HAVE A CONDITION OTHER THAN A YEAST INFECTION. CONSULT YOUR DOCTOR. If your symptoms return within two months or if you have infections that do not clear up easily with proper treatment, consult your doctor. You could be pregnant or there could be a serious underlying medical cause for your infections, including diabetes or a damaged immune system (including damage from infection with HIV—the virus that causes AIDS). (PLEASE READ ENCLOSED EDUCATIONAL PAMPHLET.) Do not use during pregnancy except under the advice and supervision of a doctor. Do not use tampons while using this medication. Keep this and all drugs out of the reach of children. In case of accidental ingestion, seek professional assistance or contact a Poison Control Center immediately. **NOT FOR USE IN CHILDREN LESS THAN 12 YEARS OF AGE.**

Active Ingredient: Clotrimazole 1%

Store at room temperature between 2° and 30°C (36° and 86°F).

Control No. & Exp. Date—See Crimp.

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TUBE LABEL